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Feb 2003

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TOXDET-03-04

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KNOWN HARMFUL EFFECTS OF CONSTITUENTS OF JET OIL CABIN SMOKE

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ABSTRACT

The construction of cabin pressurization systems of certain commercial aircraft allows pyrolyzed jet oil to leak into the cabin air, often producing visible smoke. The principal toxic constituents of this smoke are tricresyl phosphate, carbon monoxide, and N-phenyl-L-naphthylamine. Long-term neurological effects alleged by airline workers could be due to tricresyl phosphate and/or carbon monoxide exposure.

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GENERATION AND CONSTITUENTS OF JET OIL CABIN SMOKE

The BAe-146 aircraft uses precombustion air from the jet engines to pressurize the cabin. At the point at which it is bled off, the air is at a temperature of approximately 500 °C. Failing oil seals can permit leakage of jet oil into the air prior to its being bled off (1). Catalytic converters are designed to remove pollutants, but are insufficient in the event of a seal failure; smoke can often be observed in the cabin (1). A number of complaints have been made by personnel working on these commercial aircraft of long-term neurological effects. Investigations of jet oils under simulated pyrolysis conditions found tricresyl phosphates and Carbon monoxide (CO), as well as some Carbon dioxide (CO₂); trimethyl propane phosphate, Nitrogen dioxide (NO₂) and Hydrogen cyanide (HCN) were not found (1). Other potential constituents were not specifically addressed.

TRICRESYL PHOSPHATE

Tricresyl phosphate (CAS 1330-78-5) is a mixture of isomers, with the primary toxic form being the ortho isomer; formulations made in recent decades have had decreased levels of the toxic isomer (2). Studies using oral dosing suggest that current formulations have a fairly high threshold to produce an acute effect: doses of up to 2 grams per kilogram of organism did not produce any lasting neurological effects (2, 3, 4). It should be noted that route of administration can be important in the dose required to see an effect from toxic substances; no completed studies on inhalation of jet engine oils or tricresyl phosphate were available. One study which assessed the impact of orally-administered jet engine oil on enzyme activities in the brains of hens found that significant differences occurred in brain chemistry between six weeks exposure when no effect was seen, and 13 weeks, when a 23 to 34% enzyme inhibition could be detected (4). Another study, using dermal exposure in cats, found that exposure levels that did not

initially cause an effect began causing typical organophosphate poisoning effects after an initial exposure period, and that this period was itself dose-dependant (5). Therefore, doses below the ... recognized threshold for acute exposure could cause an organophosphate-type poisoning if the exposure was long in duration. Tricresyl phosphate also causes a dramatic decrease in fertility in male rats, but not in females (6).

CARBON MONOXIDE

Carbon monoxide (CO) was also produced under simulated oil leak conditions, at levels up to 100 ppm (1). Acute exposure to CO produces headache, dizziness, and nausea; long term exposure can cause memory defects and central nervous system damage, among other effects (7). Mice repeatedly exposed to CO exhibit neurodegeneration in the hippocampal region of the brain, and experience marked learning deficits (8). While most study regarding CO focuses on high exposure levels, some studies on lower exposures (0.28 to 2.8 ppm, 100 ppm) have shown a potential mechanism for oxidative damage to mammalian systems at these exposure levels: release of nitric oxide and the production of a toxic metabolite, peroxynitrite (9, 10).

N-PHENYL-1-NAPHTHYLAMINE

N-phenyl-1-naphthylamine (PAN; CAS # 90-30-2) is an antioxidant most often used in rubber formulations. It is readily absorbed by mammalian systems and rapidly converted to metabolites (11). Most toxicological studies focus on its potential carcinogenicity. One study, using both PAN and the related compound N-phenyl-2-naphthylamine administered subcutaneously to mice found a heightened incidence of lung and kidney cancers (12). A high incidence of various forms of cancer was also found among workers exposed to antirust oil containing PAN (13).

NEUROLOGICAL EFFECTS SUMMARY

Both tricresyl phosphate and CO have been suggested to cause long-term neurological complications. From the available data, the only potential neurological effect of the naphthylamines would be oncological, but there is no evidence to suggest a connection between PAN and brain tumor formation.

LITERATURE CITED

- 1. van Netten C and Leung V. 2000. Comparison of the constituents of two jet engine lubricatring oils and their volatile pyrolytic degradation products. *Appl Occup Environ Hyg* 15 (3): 277-83
- 2. Mackerer CR, Barth ML, Krueger AJ, Chawla B, and Roy TA. Comparison of neurotoxic effects and potential risks from oral administration or ingestion of tricresyl phosphate and jet engine oil containing tricresyl phosphate. J Toxicol Environ Health Part A 57 (5): 293-328
- 3. Craig PH and Barth ML. 1999. Evaluation of the hazards of industrial exposure to tricresyl phosphate: a review and interpretation of the literature. J Toxicol Environ Health Part B 2 (4): 281-300
- 4. Daughtrey W, Biles R, Jortner B, and Elrich M. 1996. Subchronic delayed neurotoxicity evaluation of jet engine lubricants containing phosphorous additives. *Fund Appl Toxicol* 32 (2): 244-249
- Abou-Donia MB, Trofatter LP, Graham DG, and Lapadul DM. 1986. Electromyographic, neuropathologic, and functional correlates in the cat as the result of tri-o-cresyl phosphate delayed neurotoxicity. *Toxicol Appl Pharmacol* 83: 126-141
- 6. Latendresse JR, Brooks CL, Flemming CD, and Capen CC. 1994. Reproductive toxicity of butylated triphenyl phosphate fluids in F344 rats. *Fundam Appl Toxicol* 223: 392-9
- 7. World Health Organization.1986. Diseases caused by asphyxiants: Carbon monoxide, Hydrogen cyanide and its toxic derivatives, and Hydrogen sulfide. Early Detection of Occupational Diseases. World Health Organization, Geneva. Pp. 154-164.
- Maurice T, Phan V, Sandillon F, and Urani A. 2000. Differential effect of dehydroepiandrosterone and its steroid precursor pregnenolone against the behavioral deficits in CO-exposed mice. Eur J Pharmacol 390: 145-55

- 9. Thom SR, Yu YA, and Ischiropoulos H. 1997. Vascular endothelial cells generate peroxynitrite in response to carbon monoxide exposure. *Chem Res Toxicol* 10: 1023-31
- Thom SR and Ischiropoulos H. 1997. Mechanism of oxidative stress from low levels of carbon monoxide. Res Rep Health Eff Inst 80: 1-19
- 11. Miyazaki K, Kawai S, Sasayama T, Iseki K, and Arita T. Absorption, metabolism and excretion of N-phenyl-1-naphthylamine in rat. *Arch Pract Pharm* 47:17-22
- 12. Wang H-W, Wang D, and Dzeng R-W. 1984. Carcinogenicity of n-phenyl-1-naphthylamine and n-phenyl-2-naphthylamine in mice. Cancer Research 44 (7): 3098-3100
- 13. Jarvholm B and Lavenius B. 1981. A cohort study on cancer among workers exposed to an antirust oil. Scand J Work Environ Health 7 (3): 179-184

REPORT DOCUMENTATION PAGE				Form Approved OMB No. 0704-0188
Public reporting burden for this collection of information is estimated to average 1 hour per response, including the time for reviewing instructions, searching existing data sources, gathering and maintaining the data needed, and completing and reviewing the collection of information. Send comments regarding this burden estimate or any other aspect of this collection of information, including suggestions for reducing this burden, to Washington Headquarters Services. Directorate for information Operations and Reports, 1215 Jefferson Davis Highway, Suite 1204, Arlington, VA 22202-4302, and to the Office of Management and Budget, Paperwork Reduction Project (0704-0188) Washington, DC 20503.				
1. AGENCY USE ONLY (Leav				
4. TITLE AND SUBTITLE Known Harmful Effects Of Constituents Of Jet Oil Cabin Smoke				5. FUNDING NUMBERS
6. AUTHOR(S) Andrew J. Bobb, Ph.D. Kenneth R. Still Ph. D. MSC USN				
7. PERFORMING ORGANIZATION NAME(S) AND ADDRESS(ES) Naval Health Research Center Detachment Toxicology				8. PERFORMING ORGANIZATION REPORT NUMBER
NHRC/TD 2612 Fifth Street, Building 433 Area B				TOXDE1-03-0-4
Wright-Patterson AFB, OH 45433-7903				
9. SPONSORING/MONITORING AGENCY NAME(S) AND ADDRESS(ES) Naval Health Research Center Detachment Toxicology NHRC/TD 2612 Fifth Street, Building 433 Area B Wright-Patterson AFB, OH 45433-7903				10. SPONSORING/MONITORING AGENCY REPORT NUMBER
11. SUPPLEMENTARY NOTES				
12a. DISTRIBUTION AVAILABILITY STATEMENT				12b. DISTRIBUTION CODE
Approved for public release; distribution is unlimited.				
13. ABSTRACT (Maximum 200 words) The construction of cabin pressurization systems of certain commercial aircraft allows pyrolyzed jet oil to leak into the cabin air, often producing visible smoke. The principal toxic constituents of this smoke are tricresyl phosphate, carbon monoxide, and N-phenyl-L-naphthylamine. Long-term neurological effects alleged by airline workers could be due to tricresyl phosphate and/or carbon monoxide exposure.				
14. SUBJECT TERMS jet oil smoke, tricresyl phosphate, N-phenyl-1-naphthylamine, carbon monoxide				15. NUMBER OF PAGES
Joe on omone, area-of phoophato, a phony: a maphinist and monomial				16. PRICE CODE
17. SECURITY CLASSIFI- CATION OF REPORT		URITY CLASSIFI- ION OF THIS PAGE	19. SECURITY CLASSIFI- CATION OF ABSTRACT	20. LIMITATION OF ABSTRACT
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